Amendments to the claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

1. (original) A compound of formula (I):

(I)

wherein

A is a fused 5-membered heteroaryl ring optionally substituted by up to two substituents independently selected from C_{1-6} alkyl, - $(CH_2)_k$ - C_{3-7} cycloalkyl, halogen, -CN, trifluoromethyl, - $(CH_2)_k$ OR 3 , - $(CH_2)_k$ CO $_2$ R 3 , - $(CH_2)_k$ NR 3 R 4 , - $(CH_2)_k$ NHCOR 3 , - $(CH_2)_k$ SO $_2$ NR 3 R 4 , - $(CH_2)_k$ NHSO $_2$ R 3 , - $(CH_2)_k$ SO $_2$ (CH $_2$) $_m$ R 5 , a 5- or 6-membered heterocyclyl ring containing nitrogen optionally substituted by C_{1-2} alkyl or - $(CH_2)_k$ CO $_2$ R 3 , and a 5-membered heteroaryl ring optionally substituted by C_{1-2} alkyl;

A is a fused 5-membered heteroaryl ring substituted by $-BR^6$, and A is optionally further substituted by one substituent selected from $-OR^7$, halogen, trifluoromethyl, -CN, $-CO_2R^7$ and C_{1-6} alkyl optionally substituted by hydroxy;

A is a fused 5-membered heteroaryl ring substituted by - $(CH_2)_n$ heterocyclyl wherein the heterocyclyl is a 5- or 6-membered heterocyclic ring containing one or two heteroatoms independently selected from oxygen, sulfur and nitrogen optionally substituted by up to two substituents independently selected from oxo, C_{1-6} alkyl, - $(CH_2)_p$ phenyl, - OR^7 , - $(CH_2)_p$ CO₂R⁷, - NR^7 R⁸ and - $CONR^7$ R⁸, and A is optionally further substituted by one substituent selected from - OR^7 , halogen, trifluoromethyl, -CN, - CO_2 R⁷ and C_{1-6} alkyl optionally substituted by hydroxy; or

A is a fused 5-membered heteroaryl ring substituted by - $(CH_2)_q$ aryl or - $(CH_2)_q$ heteroaryl wherein the aryl or heteroaryl is optionally substituted by one or more substituents independently selected from oxo, C_{1-6} alkyl, halogen, -CN,

trifluoromethyl, -OR 9 , -(CH $_2$) $_r$ CO $_2$ R 10 , -NR 9 R 10 , -(CH $_2$) $_r$ CONR 9 R 10 , -NHCOR 9 , -SO $_2$ NR 9 R 10 , -NHSO $_2$ R 9 and -S(O) $_s$ R 9 , and

A is optionally further substituted by one substituent selected from -OR⁷, halogen, trifluoromethyl, -CN, -CO₂R⁷ and C_{1-6} alkyl optionally substituted by hydroxy;

R¹ is selected from methyl and chloro;

R² is selected from -NH-CO-R¹¹ and -CO-NH-(CH₂)_t-R¹²;

 R^3 is selected from hydrogen, $C_{1\text{-}6}$ alkyl optionally substituted by up to two OH groups, -(CH₂)_k-C₃₋₇cycloalkyl, -(CH₂)_kphenyl optionally substituted by R^{13} and/or R^{14} and -(CH₂)_kheteroaryl optionally substituted by R^{13} and/or R^{14} ,

 R^4 is selected from hydrogen and C_{1-6} alkyl, or

 R^3 and R^4 , together with the nitrogen atom to which they are bound, form a 5-or 6-membered heterocyclic ring optionally containing one additional heteroatom selected from oxygen, sulfur and N-R¹⁵;

 R^5 is selected from $C_{1\text{-}6}$ alkyl optionally substituted by up to three halogen atoms, $C_{2\text{-}6}$ alkenyl optionally substituted by phenyl, $C_{3\text{-}7}$ cycloalkyl, heteroaryl optionally substituted by up to three R^{13} and/or R^{14} groups, and phenyl optionally substituted by R^{13} and/or R^{14} ;

 R^6 is a C_{3-6} alkyl group substituted by at least two substituents independently selected from -OR¹⁶, -NR¹⁶R¹⁷, -CO₂R¹⁶, -CONR¹⁶R¹⁷, -NHCOR¹⁶ and -NHSO₂R¹⁶;

 ${\rm R}^7$ and ${\rm R}^8$ are each independently selected from hydrogen and ${\rm C}_{1\text{-}6}$ alkyl;

 R^9 is selected from hydrogen, -(CH₂)_u-C₃₋₇cycloalkyl, -(CH₂)_uheterocyclyl, -(CH₂)_uaryl, and C₁₋₆alkyl optionally substituted by up to two substituents independently selected from -OR 18 and -NR 18 R 19 ,

R¹⁰ is selected from hydrogen and C₁₋₆alkyl, or

 R^9 and R^{10} , together with the nitrogen atom to which they are bound, form a 5- or 6-membered heterocyclic ring optionally containing one additional heteroatom selected from oxygen, sulfur and N-R¹⁵;

 R^{11} is selected from hydrogen, C₁₋₆alkyl, -(CH₂)_t-C₃₋₇cycloalkyl, trifluoromethyl, -(CH₂)_vheteroaryl optionally substituted by R^{20} and/or R^{21} , and -(CH₂)_vphenyl optionally substituted by R^{20} and/or R^{21} ;

 R^{12} is selected from hydrogen, $C_{1\text{-}6}$ alkyl, $C_{3\text{-}7}$ cycloalkyl, -CONHR²², phenyl optionally substituted by R^{20} and/or R^{21} , and heteroaryl optionally substituted by R^{20} and/or R^{21} ;

 R^{13} and R^{14} are each independently selected from halogen, -CN, trifluoromethyl, nitro, C1-6alkyl, C1-6alkoxy, -CONR^22R^23, -COR^24, -CO_2R^24, and heteroaryl, or

 R^{13} and R^{14} are linked to form a fused 5-membered heterocyclyl ring containing one heteroatom selected from oxygen, sulfur and N-R¹⁵, or a fused heteroaryl ring;

R¹⁵ is selected from hydrogen and methyl;

 $R^{16},\,R^{17},\,R^{18}$ and R^{19} are each independently selected from hydrogen and $C_{1\text{-}6}alkyl;$

 R^{20} is selected from $C_{1\text{-}6}$ alkyl, $C_{1\text{-}6}$ alkoxy, -(CH $_2$)t-C $_3$ -7cycloalkyl, - CONR 22 R 23 , -NHCOR 23 , halogen, -CN, -(CH $_2$)wNR 25 R 26 , trifluoromethyl, phenyl optionally substituted by one or more R^{21} groups, and heteroaryl optionally substituted by one or more R^{21} groups;

 R^{21} is selected from C_{1-6} alkyl, C_{1-6} alkoxy, halogen, trifluoromethyl, and $-(CH_2)_wNR^{25}R^{26}$;

 R^{22} and R^{23} are each independently selected from hydrogen and $C_{1\text{-}6}$ alkyl, or R^{22} and R^{23} , together with the nitrogen atom to which they are bound, form a 5- or 6-membered heterocyclic ring optionally containing one additional heteroatom selected from oxygen, sulfur and N-R 15 , wherein the ring may be substituted by up to two $C_{1\text{-}6}$ alkyl groups;

R²⁴ is C₁₋₆alkyl;

 R^{25} is selected from hydrogen, $C_{1\text{-}6}$ alkyl and -(CH₂)_t-C₃₋₇cycloalkyl optionally substituted by $C_{1\text{-}6}$ alkyl,

R²⁶ is selected from hydrogen and C₁₋₆alkyl, or

 R^{25} and R^{26} , together with the nitrogen atom to which they are bound, form a 5- or 6-membered heterocyclic ring optionally containing one additional heteroatom selected from oxygen, sulfur and $N-R^{15}$;

R²⁷ is hydrogen or C₁₋₆alkyl;

B is selected from a bond, oxygen, NH and $S(O)_X$;

X and Y are each independently selected from hydrogen, methyl and halogen;

Z is selected from halogen, C_{1-6} alkyl and -OR²⁷;

k, m and w are each independently selected from 0, 1, 2 and 3;

n, q, r, s, t and x are each independently selected from 0, 1 and 2; and u and v are each independently selected from 0 and 1;

or a pharmaceutically acceptable derivative thereof.

2. (original) A compound according to claim 1 wherein A is a fused 5-membered heteroaryl ring containing up to two heteroatoms independently selected from oxygen and nitrogen.

International Application No. PCT/GB2005/000266
International Filing Date: 27 January 2005

- 3. (currently amended) A compound according to claim 1 or claim 2 wherein A is substituted by -(CH₂)_qaryl or -(CH₂)_qheteroaryl wherein the aryl or heteroaryl is optionally substituted by one or more substituents independently selected from oxo, C_{1-6} alkyl, halogen, -CN, trifluoromethyl, -OR⁹, -(CH₂)_rCO₂R¹⁰, -NR⁹R¹⁰, -(CH₂)_rCONR⁹R¹⁰, -NHCOR⁹, -SO₂NR⁹R¹⁰, -NHSO₂R⁹ and -S(O)_sR⁹.
- 4. (currently amended) A compound according to <u>claim 1</u> any one of the <u>preceding claims</u> wherein R¹ is methyl.
- 5. (currently amended) A compound according to <u>claim 1 any one of the preceding claims</u>-wherein R² is -CO-NH-(CH₂)_t-R¹².
- 6. (currently amended) A compound according to <u>claim 1 any one of the preceding claims</u> wherein X is hydrogen or fluorine.
- 7. (original) A compound according to claim 1 substantially as hereinbefore defined with reference to any one of Examples 1 to 6, or a pharmaceutically acceptable derivative thereof.
- 8. (original) A compound selected from:

N-cyclopropyl-3-[5-fluoro-3-(4-pyridinyl)-1H-indazol-6-yl]-4-methylbenzamide; and N-cyclopropyl-3-fluoro-5-[5-fluoro-3-(4-pyridinyl)-1,2-benzisoxazol-6-yl]-4-methylbenzamide;

or a pharmaceutically acceptable derivative thereof.

- 9. (currently amended) A pharmaceutical composition comprising at least one compound as claimed in <u>claim 1 any one of claims 1 to 8</u>, or a pharmaceutically acceptable derivative thereof, in association with one or more pharmaceutically acceptable excipients, diluents and/or carriers.
- 10. (currently amended) A compound according to <u>claim 1 any one of claims 1</u> to 8, or a pharmaceutically acceptable derivative thereof, for use in therapy.
- 11. (currently amended) A compound as claimed in <u>claim 1 any one of claims 1 to 8</u>, or a pharmaceutically acceptable derivative thereof, for use in the treatment or prophylaxis of a condition or disease state mediated by p38 kinase activity or mediated by cytokines produced by the activity of p38 kinase.

12. (currently amended) A method for treating a condition or disease state mediated by p38 kinase activity or mediated by cytokines produced by the activity of p38 kinase comprising administering to a patient in need thereof a compound as claimed in claim 1 any one of claims 1 to 8, or a pharmaceutically acceptable derivative thereof.

13. (cancelled)

14. (currently amended) A process for preparing a compound of formula (I) as claimed in <u>claim 1</u> any one of claims 1 to 8, or a pharmaceutically acceptable derivative thereof, which comprises

(a) reacting a compound of formula (II)

(II)

in which A is defined in claim 1 and Hal is halogen, with a compound of formula (IIIA) or (IIIB)

(IIIA)

(IIIB)

in which R^1 , R^2 , X and Y are as defined in claim 1, in the presence of a catalyst, or

- (b) final stage modification of one compound of formula (I) as defined in claim 1 to give another compound of formula (I) as defined in claim 1.
- 15. (new) A compound according to claim 2 wherein A is substituted by $-(CH_2)_q \text{aryl or } -(CH_2)_q \text{heteroaryl wherein the aryl or heteroaryl is optionally substituted by one or more substituents independently selected from oxo, C_{1-6} alkyl, halogen, -CN, trifluoromethyl, -OR$^9, -(CH_2)_r CO_2 R^{10}, -NR^9 R^{10}, -(CH_2)_r CO_2 R^{10}, -NR^9 R^{10}, -(CH_2)_r CO_2 R^{10}, -NHSO_2 R^9 \text{ and } -S(O)_s R^9.$
- 16. (new) A compound according to claim 15 wherein \mathbb{R}^1 is methyl.
- 17. (new) A compound according to claim 15 wherein R² is -CO-NH-(CH₂)_t-R¹².
- 18. (new) A compound according to claim 15 wherein X is hydrogen or fluorine.